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=> fil reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 6 SEP 2002 HIGHEST RN 447682-31-7 DICTIONARY FILE UPDATES: 6 SEP 2002 HIGHEST RN 447682-31-7

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

PSEUDOMESON/BI

```
=> e pseudomanas
E1
           1
                  PSEUDOMALACHITE/BI
E2
          105
                  PSEUDOMALLEI/BI
E3
            2 --> PSEUDOMANAS/BI
                 PSEUDOMANIBACANINE/BI
E4
            1
                 PSEUDOMECONIN/BI
E5
            1
                 PSEUDOMENOPON/BI
E6
            2
                 PSEUDOMERU/BI
            5
E7
                 PSEUDOMERUCATH/BI
E8
            5
                 PSEUDOMERUCATHIN/BI
E9
            2
                 PSEUDOMERUCATHINE/BI
E10
            5
E11
            3
                  PSEUDOMESENTEROIDES/BI
```

=> s e3

E12

L1 2 PSEUDOMANAS/BI

=> e streptococcus

El	1		STREPTOCOCCI/BI
E2	8		STREPTOCOCCIN/BI
E3	38201 -	>	STREPTOCOCCUS/BI
E4	1		STREPTOCOM/BI
E5	1		STREPTOCYANIN/BI
E6	1		STREPTOCYANINE/BI
E7	1		STREPTOCYCLIN/BI
E8	3		STREPTOCYCLINE/BI
E9	2		STREPTOCYMES/BI
E10	1		STREPTODECA/BI
E11	1		STREPTODECASE/BI
E12	4		STREPTODORN/BI

=> s e3

L2 38201 STREPTOCOCCUS/BI

=> e streptococcus/cn

E1 1 STREPTOCOCCIN A-M49 (STREPTOCOCCUS PYOGENES STRAIN GT9538 GE NE SCNA'' PRECURSOR)/CN

E2 1 STREPTOCOCCIN SAL-P/CN

```
0 --> STREPTOCOCCUS/CN
E3
                   STREPTOCOCCUS AGALACTIAE/CN
E4
             1
                   STREPTOCOCCUS CREMORIS SERINE PROTEINASE/CN
E5
                  STREPTOCOCCUS DIACETILACTIS NEUTRAL PROTEINASE/CN
E6
                  STREPTOCOCCUS DIACETILACTIS, CULTURE DISTILLATE FLAVORING/CN
E7
                  STREPTOCOCCUS EQUISIMILIS/CN
E8
            1
                  STREPTOCOCCUS FAECIENS/CN
E9
            1
                  STREPTOCOCCUS FAECIUM/CN
E10
            1
                  STREPTOCOCCUS LACTIS ACID PROTEINASE/CN
E11
             1
                  STREPTOCOCCUS LACTIS PROTEINASE/CN
E12
            1
=> e staphylococcus
                 STAPHYLOCOCCCUS/BI
            6
E1
E2
            16
                  STAPHYLOCOCCIN/BI
E3
         25045 --> STAPHYLOCOCCUS/BI
E4
            3
                  STAPHYLOCYSTIS/BI
E5
             2
                   STAPHYLOFERRI/BI
E6
             2
                   STAPHYLOFERRIN/BI
E7
           38
                   STAPHYLOID/BI
E8
           45
                   STAPHYLOKINASE/BI
E9
           12
                   STAPHYLOL/BI
E10
            2
                   STAPHYLOLYSIN/BI
           10
E11
                   STAPHYLOLYTICUS/BI
E12
           11
                  STAPHYLOMYCIN/BI
=> s e3
         25045 STAPHYLOCOCCUS/BI
=> e clostridium
                  CLOSTRIDIOPEPTID/BI
                  CLOSTRIDIOPEPTIDASE/BI
E2
E3
          9155 --> CLOSTRIDIUM/BI
                  CLOSTRIDIUMB24/BI
E4
            1
E5
            1
                  CLOSTRIDIUMB33/BI
E6
            1
                  CLOSTRIDIUMB40/BI
E7
            3
                  CLOSTRIDUM/BI
           10
E8
                  CLOSTRIPAIN/BI
E9
            2
                  CLOSULAM/BI
           1
E10
                  CLOSURE/BI
            1
E11
                  CLOSYL/BI
E12
                  CLOSYLATE/BI
=> s e3
          9155 CLOSTRIDIUM/BI
=> fil .search
                                                 SINCE FILE TOTAL SESSION
COST IN U.S. DOLLARS
FULL ESTIMATED COST
                                                      16.76
                                                                 16.97
FILE 'MEDLINE' ENTERED AT 14:23:55 ON 09 SEP 2002
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FILE 'USPATFULL' ENTERED AT 14:23:55 ON 09 SEP 2002
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=> d his

(FILE 'HOME' ENTERED AT 14:22:38 ON 09 SEP 2002)

FILE 'REGISTRY' ENTERED AT 14:22:45 ON 09 SEP 2002

E PSEUDOMANAS

L12 S E3

E STREPTOCOCCUS

38201 S E3 L2

E STREPTOCOCCUS/CN

E STAPHYLOCOCCUS

L3 25045 S E3

E CLOSTRIDIUM

L49155 S E3

> FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL, EMBASE' ENTERED AT 14:23:55 ON 09 SEP 2002

=> s l1 or l2 or l3 or l4

TOO MANY TERMS FOR FILE CROSSOVER IN L2

There are limits on the size of an answer set being crossed over from one file to another. Enter HELP CROSSOVER at an arrow prompt (=>) for specific information.

=> fil reg

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 3.93 20.90

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 6 SEP 2002 HIGHEST RN 447682-31-7 DICTIONARY FILE UPDATES: 6 SEP 2002 HIGHEST RN 447682-31-7

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:

http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> e hemophilus influenza

E1	5	HEMOPHILIC/BI
E2	18	HEMOPHILUS/BI

E3 0 --> HEMOPHILUS INFLUENZA/BI

E4 2 HEMOPHORE/BI E5 HEMOPLAST/BI 1 E6 1 HEMOPLEX/BI

E7 3216 HEMOPOIETIC/BI E8 103 HEMOPOIETIN/BI

E9 HEMOPOIETINS/BI 2

E10 HEMOPR/BI

```
HEMOPRENE/BI
             1
E11
                    HEMOPROTECTOR/BI
E12
             5
=> s e2
            18 HEMOPHILUS/BI
=> e streptococus pneumoniae
                   STREPTOCOCCIN/BI
E2
         38201
                   STREPTOCOCCUS/BI
E3
             0 --> STREPTOCOCUS PNEUMONIAE/BI
E4
             1
                   STREPTOCOM/BI
E5
             1
                   STREPTOCYANIN/BI
E6
             1
                   STREPTOCYANINE/BI
E7
             1
                   STREPTOCYCLIN/BI
E8
                   STREPTOCYCLINE/BI
E9
             2
                   STREPTOCYMES/BI
E10
                   STREPTODECA/BI
             1
E11
                   STREPTODECASE/BI
             1
E12
                   STREPTODORN/BI
=> e streptococus?
                   STREPTOCOCCIN/BI
E2
         38201
                   STREPTOCOCCUS/BI
E3
             0 --> STREPTOCOCUS?/BI
E4
                   STREPTOCOM/BI
             1
                   STREPTOCYANIN/BI
E5
             1
E6
             1
                   STREPTOCYANINE/BI
                   STREPTOCYCLIN/BI
E7
             1
                   STREPTOCYCLINE/BI
E8
             3
E9
             2
                   STREPTOCYMES/BI
E10
             1
                   STREPTODECA/BI
E11
             1
                   STREPTODECASE/BI
                   STREPTODORN/BI
E12
=> e streptococuspneu?
             8
                   STREPTOCOCCIN/BI
E2
         38201
                   STREPTOCOCCUS/BI
E3
             0 --> STREPTOCOCUSPNEU?/BI
E4
             1
                   STREPTOCOM/BI
E5
                   STREPTOCYANIN/BI
             1
E6
                   STREPTOCYANINE/BI
             1
E7
                   STREPTOCYCLIN/BI
             1
E8
                   STREPTOCYCLINE/BI
             3
E9
             2
                   STREPTOCYMES/BI
E10
                   STREPTODECA/BI
             1
E11
                   STREPTODECASE/BI
             1
E12
             4
                   STREPTODORN/BI
=> e streptococus pneu?
E1
             8
                   STREPTOCOCCIN/BI
E2
         38201
                  STREPTOCOCCUS/BI
E3
             0 --> STREPTOCOCUS PNEU?/BI
E4
                   STREPTOCOM/BI
             1
E5
                   STREPTOCYANIN/BI
             1
E6
             1
                   STREPTOCYANINE/BI
E7
             1
                   STREPTOCYCLIN/BI
E8
             3
                   STREPTOCYCLINE/BI
E9
             2
                   STREPTOCYMES/BI
E10
             1
                   STREPTODECA/BI
E11
             1
                   STREPTODECASE/BI
E12
             4
                   STREPTODORN/BI
```

```
PNEUMONANTHOSIDE/BI
           2
E1
           80
                 PNEUMONIA/BI
E2
         20452 --> PNEUMONIAE/BI
E3
                 PNEUMOPHILA/BI
E4
         475
           18
                  PNEUMOPHILIA/BI
E5
                  PNEUMORA/BI
E6
            3
                  PNEUMOS/BI
E7
            4
E8
            4
                  PNEUMOSAMINE/BI
E9
            2
                  PNEUMOTROPICA/BI
E10
           99
                  PNEUMOVIRUS/BI
E11
            1
                  PNEUMOXIDE/BI
E12
           62
                  PNEUSTES/BI
=> s e3
L6
        20452 PNEUMONIAE/BI
=> s 16 and12
MISSING OPERATOR
=> s 16 and 12
       12092 L6 AND L2
=> e fasciae
E1
         168
                  FASCI/BI
E2
            3
                  FASCIA/BI
E3
            0 --> FASCIAE/BI
               FASCIANS/BI
E4
           31
E5
           96
                 FASCIATA3/BI
E6
            1
                 FASCIATA4/BI
E7
            1
                  FASCIATA5/BI
E8
            1
E9
            1
                 FASCIATIN/BI
                 FASCIATO/BI
E10
            1
E11
            1
                 FASCIATOXIN/BI
E12
            9
                  FASCIATUM/BI
=> s e2
L8
            3 FASCIA/BI
=> s 18 and 12
L9
           0 L8 AND L2
=> e listeria
E1
           6
                 LISTERA/BI
                 LISTERI/BI
E2
           13
E3
        10507 --> LISTERIA/BI
E4
                 LISTERIAL/BI
           1
E5
            2
                 LISTERINE/BI
E6
            1
                  LISTERIOCIN/BI
           10
E7
                  LISTERIOL/BI
E8
            9
                  LISTERIOLYSIN/BI
                  LISTERIOLYSINS/BI
E9
            3
           1
E10
                  LISTERITE/BI
E11
           3
                  LISTEROL/BI
E12
            2
                  LISTEROLYSIN/BI
=> s e3
L10
        10507 LISTERIA/BI
=> e salmonella
E1
     5
                 SALMONASE/BI
E2
        14768
                SALMONE/BI
E3
        14766 --> SALMONELLA/BI
```

```
1
                    SALMONEUM/BI
E4
            22
                    SALMONIC/BI
E5
E6
           115
                    SALMONICIDA/BI
            22
                    SALMONICOL/BI
E7
            22
                    SALMONICOLOR/BI
E8
                    SALMONID/BI
E9
            6
            47
                    SALMONINARUM/BI
E10
E11
            73
                    SALMONIS/BI
                    SALMONNELLA/BI
E12
             1
=> s e3
         14766 SALMONELLA/BI
L11
=> e ecoli
E1
             1
                    ECOLAN/BI
E2
             1
                   ECOLCRAFT/BI
E3
             2 --> ECOLI/BI
E4
             1
                   ECOLID/BI
E5
             1
                   ECOLIFE/BI
E6
             3
                   ECOLINE/BI
E7
             2
                   ECOLITE/BI
E8
             1
                   ECOLL/BI
E9
             1
                   ECOLL1/BI
E10
             1
                   ECOLL10/BI
E11
                   ECOLL2/BI
             1
E12
                   ECOLL3/BI
=> s e3
             2 ECOLI/BI
L12
=> e campylobacter
E1
             2
                    CAMPYLENCHIA/BI
E2
             3
                    CAMPYLIUM/BI
          2847 --> CAMPYLOBACTER/BI
E3
E4
            1
                   CAMPYLOCARPUM/BI
E5
             6
                   CAMPYLOCENTRUS/BI
E6
            26
                   CAMPYLOMORMYRUS/BI
E7
             2
                   CAMPYLOPH/BI
             2
                   CAMPYLOPHYL/BI
E8
            2
E9
                   CAMPYLOPHYLLA/BI
                   CAMPYLOPODUM/BI
E10
            1
E11
            19
                   CAMPYLOPUS/BI
                   CAMPYLOPUSAUR/BI
E12
            1
=> s e3
L13
          2847 CAMPYLOBACTER/BI
=> e streptococcus(w) mutans
E1
             8
                 STREPTOCOCCIN/BI
         38201
E2
                   STREPTOCOCCUS/BI
E3
             0 --> STREPTOCOCCUS (W) MUTANS/BI
E4
                   STREPTOCOM/BI
             1
E5
             1
                   STREPTOCYANIN/BI
E6
             1
                   STREPTOCYANINE/BI
E7
             1
                   STREPTOCYCLIN/BI
E8
             3
                   STREPTOCYCLINE/BI
E9
             2
                   STREPTOCYMES/BI
E10
             1
                   STREPTODECA/BI
E11
             1
                   STREPTODECASE/BI
E12
                   STREPTODORN/BI
=> e mutans
                   MUTANOL/BI
E1
             1
```

MUTANOLYSIN/BI E2 1 430 --> MUTANS/BI E3 MUTANSUCR/BI E41 MUTANSUCRASE/BI E5 1 4858 MUTANT/BI E6 MUTANTS/BI E7 30 MUTAROT/BI E8 16 E9 16 MUTAROTASE/BI E10 1 MUTAS/BI E11 1177 MUTASE/BI E12 1 MUTASES/BI => s e3 430 MUTANS/BI L14 => s 12 and 114 425 L2 AND L14 L15 => s mycobacterium 12105 MYCOBACTERIUM 2 MYCOBACTERIA L16 12107 MYCOBACTERIUM (MYCOBACTERIUM OR MYCOBACTERIA) => fil .search COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 38.66 59.56 FILE 'MEDLINE' ENTERED AT 14:28:38 ON 09 SEP 2002 FILE 'CAPLUS' ENTERED AT 14:28:38 ON 09 SEP 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'BIOSIS' ENTERED AT 14:28:38 ON 09 SEP 2002 COPYRIGHT (C) 2002 BIOLOGICAL ABSTRACTS INC. (R) FILE 'USPATFULL' ENTERED AT 14:28:38 ON 09 SEP 2002 CA INDEXING COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'EMBASE' ENTERED AT 14:28:38 ON 09 SEP 2002 COPYRIGHT (C) 2002 Elsevier Science B.V. All rights reserved. => d his (FILE 'HOME' ENTERED AT 14:22:38 ON 09 SEP 2002) FILE 'REGISTRY' ENTERED AT 14:22:45 ON 09 SEP 2002 E PSEUDOMANAS L12 S E3 E STREPTOCOCCUS L238201 S E3 E STREPTOCOCCUS/CN E STAPHYLOCOCCUS

FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL, EMBASE' ENTERED AT 14:23:55 ON 09 SEP 2002

L3

L4

25045 S E3

9155 S E3

E CLOSTRIDIUM

```
FILE 'REGISTRY' ENTERED AT 14:24:14 ON 09 SEP 2002
                E HEMOPHILUS INFLUENZA
L5
             18 S E2
                E STREPTOCOCUS PNEUMONIAE
                E STREPTOCOCUS?
                E STREPTOCOCUSPNEU?
                E STREPTOCOCUS PNEU?
                E PNEUMONIAE
L<sub>6</sub>
          20452 S E3
L7
          12092 S L6 AND L2
                E FASCIAE
              3 S E2
L8
              0 S L8 AND L2
L9
                E LISTERIA
          10507 S E3
L10
                E SALMONELLA
          14766 S E3
L11
                E ECOLI
              2 S E3
L12
                E CAMPYLOBACTER
L13
           2847 S E3
                E STREPTOCOCCUS (W) MUTANS
                E MUTANS
L14
            430 S E3
L15
            425 S L2 AND L14
L16
          12107 S MYCOBACTERIUM
     FILE 'MEDLINE, CAPLUS, BIOSIS, USPATFULL, EMBASE' ENTERED AT 14:28:38 ON
     09 SEP 2002
=> s l1 or l3 or l4 or l5 or l6 or l7 or l8 or l9lr l10 or l11 or l12 or l13 or l14
or 115 or 116
MISSING OPERATOR L9LR L10
The search profile that was entered contains terms or
nested terms that are not separated by a logical operator.
=> s 11 or 13 or 14 or 15 or 16 or 17 or 18 or 19 or 110 or 111 or 112 or 113 or
114 or 115 or 116
TOO MANY TERMS FOR FILE CROSSOVER IN L3
There are limits on the size of an answer set being crossed over from
one file to another. Enter HELP CROSSOVER at an arrow prompt (=>)
for specific information.
=> s l1 or l4 or l5 or l6 or l7 or l8 or l9 or l10 or l11 or l12 or l13 or l14 or
115 or 116
TOO MANY TERMS FOR FILE CROSSOVER IN L6
There are limits on the size of an answer set being crossed over from
one file to another. Enter HELP CROSSOVER at an arrow prompt (=>)
for specific information.
=> s l1 or l4 or l5 or l7 or l8 or l9 or l10 or l11 or l12 or l13 or l14 or l15 or
TOO MANY TERMS FOR FILE CROSSOVER IN L7
There are limits on the size of an answer set being crossed over from
one file to another. Enter HELP CROSSOVER at an arrow prompt (=>)
for specific information.
=> s l1 or l4 or 15 or 18 or 19 or 110 or 111 or 112 or 113 or 114 or 115 or 116
TOO MANY TERMS FOR FILE CROSSOVER IN L10
There are limits on the size of an answer set being crossed over from
one file to another. Enter HELP CROSSOVER at an arrow prompt (=>)
for specific information.
```

=> s l1 or l4 or l5 or l8 or l9 or l11 or l12 or l13 or l14 or l15 or l16 TOO MANY TERMS FOR FILE CROSSOVER IN L11 There are limits on the size of an answer set being crossed over from one file to another. Enter HELP CROSSOVER at an arrow prompt (=>) for specific information.

=> s l1 or l4 or l5 or l8 or l9 or l12 or l13 or l14 or l15 or l16 TOO MANY TERMS FOR FILE CROSSOVER IN L16 There are limits on the size of an answer set being crossed over from one file to another. Enter HELP CROSSOVER at an arrow prompt (=>) for specific information.

=> s l1 or l4 or l5 or l8 or l9 or l12 or l13 or l14 or l15
1 FILES SEARCHED...
3 FILES SEARCHED...
4 FILES SEARCHED...

L17 25859 L1 OR L4 OR L5 OR L8 OR L9 OR L12 OR L13 OR L14 OR L15

=> s l17 and (lytic(w)enzyme? or lyze?)
L18 29 L17 AND (LYTIC(W) ENZYME? OR LYZE?)

=> dup rem 119
PROCESSING COMPLETED FOR L19
L20 4 DUP REM L19 (0 DUPLICATES REMOVED)

=> d ibib ab 1-YOU HAVE REQUESTED DATA FROM 4 ANSWERS - CONTINUE? Y/(N):y

```
NUMBER OF CLAIMS: 11

REMPMIARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Page(s)

LINE COUNT: 742

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Methods to obtain genetic modifications of cells in histoculture are described. Modification is assisted by treating the histoculture with collagenase prior to contacting the histoculture with the delivery vehicle for the desired gene. Heir follicles and other organized tissues
                                  can be modified in this was and then transplanted into intact recipients.
    tissues
   L20 ANSWER 3 OF 4 USPATPULL ACCESSION NUMBER: 90:50 TITLE: Metho
                                                                                                               ATPULL
90:50741 USPATFULL
Methods for separating malignant cells from clinical
specimens
Rotman, M. Boris, Jamestown, RI, United States
Brown University Research Foundation, Providence, RI,
United States (U.S. corporation)
    INVENTOR(S):
PATENT ASSIGNEE(S):
                                                                                                                NUMBER KIND DATE

US 4937187 19900626
US 1987-11219 19870205 (7)
Continuation-in-part of Ser. No. US 1984-623183, filed on 22 Jun 1984, now patented, Pat. No. US 4734372
    PATENT INFORMATION:
APPLICATION INFO.:
RELATED APPLN. INFO.:
                                                                                                                is a continuation-in-part of Ser. No. US 1983-463669, filed on 4 Feb 1983, now patented, Pat. No. US 4559299 Utility Granted Rosen, Sam Engellenner, Thomas J. 19
is a continuation-in-part of Ser. No. US 1983-463689,
filed on 4 Feb 1983, now patented, Pat. No. US 4559299
DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINE: Rosen, Sam
LEGAL REPRESENTATIVE: Engellenner, Thomas J.

NUMBER OF CLAIMS: 19
EXEMPLARY CIAIM: 115
LINE COUNT: 1 415
CAS INDEXING 15 AVAILABLE FOR THIS PATENT.
AB Pragments of a biopsy asmple on the order of about 50 to 5000 cells are preferred for establishing viable tumor cell cultures for purposes such as establishing cell lines, chemotherapeutic assays and the like. Such fragments retain the three-dimensional cellular structure or organization of the original tumor and, therefore, can be cultured more readily. To obtain such fragments suitable for culturing, the biopsy sample can be enzymatically dispated in a proteolytic or nucleolytic enzyme, such as collagenase, or by mechanical dissociation, or both where necessary. The fragments can then be suspended in an aqueous medium so that non-aggregated cells (e.g., red blood cells, lymphocytes, macrophages) and cellular debris will form a supernatant while the remaining fragments containing aggregated tumor cells are deposited in
                                   yes,
macrophages) and cellular debris will form a supernatant while the
remaining fragments containing aggregated tumor cells are deposited in
                                  sediment layer. Preferably, the medium is an isotonic tissue culture medium and decantation is conducted at least twice; first in a serum-containing medium and then, secondly, in a serum-free medium. Pragments containing living tumor cells can be selected by fluorochromasia, that is, by contacting the sedimented layer with a fluorogenic substrate such that viable tumor cells take up and
  hydrolyse the substrate, and then exhibit fluorescence. Cytotoxicity assay protocols employing tumor cell aggregates prepared by the present techniques are also disclosed.
```

L20 ANSMER 1 OF 4 USPATFULL
ACCESSION NUMBER: 2002:21826 USPATFULL
Hethods for introducing genes into mammalian subjects
Saito, Norimitau, Kanagawa, JAPAN
Zhao, Ming, San Diego, CA, UNITED STATES .

US 2002012661 US 2000-734786

PATENT INFORMATION: APPLICATION INFO.:

PRIORITY INFORMATION: DOCUMENT TYPE: FILE SEGMENT: LEGAL REPRESENTATIVE:

NUMBER KIND DATE

US 2002012661 A1 20020131
US 2000-734786 A1 20001211 (9)

US 1999-170166P 19991210 (60)
UL:11ty
APPLICATION
Kate H. Murashige, Morrison & Foerster LLP, Suite 500, 3611 Valley Centre Drive, San Diego, CA, 92130-2332
21

NUMBER DATE

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L20 ANSWER 2 OF 4 USPATFULL ACCESSION NUMBER: 2002:5
                                                                                                                                                                                        2002:9855 USPATFULL
Peptide-lipid conjugates, liposomes and lipsomal drug
delivery
Meeres, Paul R., Princeton, NJ, United States
Pak, Charles, Princeton, NJ, United States
Ali, Shaukat, Monmouth Junction, NJ, United States
Janoff, Andrew, Yardley, PA, United States
Franklin, J. Craig, Skillman, NJ, United States
Franklin, Ravi K., Plainsboro, NJ, United States
Cabral-Lilly, Donna, Princeton, NJ, United States
Ahl, Patrick L., Princeton, NJ, United States
Elan PharmaceuticalsTechnologies, Inc., King of
Prussia, PA, United States (U.S. corporation)
                                                                                                                                                                                            2002:9855 USPATFULL
   INVENTOR(S):
     PATENT ASSIGNEE (S):
                                                                                                                                                                                        NUMBER KIND DATE

US 6319069 B1 20020115
US 1999-343650 19990629 (9)
Continuation-in-part of Ser. No. US 1998-168010, filed on 7 Oct 1998, now patented, Pat. No. US 6143716 Division of Ser. No. US 1997-950618, filed on 15 Oct 1997, now patented, Pat. No. US 6087325
     PATENT INFORMATION:
APPLICATION INFO.:
RELATED APPLN. INFO.:
                                                                                                                                                                                                                                                NIMBER
                                                                                                                                                                                                                                                                                                                                                        DATE
                                                                                                                                                                                      US 1996-27544P 19961015 (60) US 1997-39183P 19970227 (60) Utility GRANTED Nguyen, Dave T. Burns, Doane, Swecker & Mathis L.L.P. 22
     PRIORITY INFORMATION:
DOCUMENT TYPE: US 1997-39183P 19970227 (60)

DOCUMENT TYPE: US 1997-39183P 19970227 (60)

DOCUMENT TYPE: US 1997-39183P 19970227 (60)

PRIMARY EXAMINER: Negree Search Sea
   conditions,
e.g., tumors, microbial infaction and inflammations,
characterized by the occurrence of peptidase-secreting cells.
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20 ANSWER 4 OF 4 USPATFULL
CCESSION NUMBER: 87:36
                                                                                                                  ATFULL
87:36077 USPATFULL
Inhibition of mammalian collagenolytic enzymes by tetracyclines
Golub, Lorne M., Smithtown, NY, United States
McNamara, Thomas P., Port Jefferson, NY, United States
Remamurthy, N. S., Smithtown, NY, United States
Research Poundation of State University, Albany, NY,
United States (U.S. corporation)
 ACCL...
  INVENTOR (S):
  PATENT ASSIGNEE(S):
                                                                                                                                                                         R KIND
                                                                                                                                            NUMBER
                                                                                                                                                                                                                                                   DATE
                                                                                                                 US 4666897
US 1983-566517
Utility
Granted
Meyera, Albert T.
Kilcoyne, John M.
Behr, Omri M.
10
NOMBER KIND DATE

PATENT INFORMATION: US 4666897 19870519

APPLICATION INFO: US 1983-566517 19831229 (6)

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

PRIMARY EXAMINER: Meyers, Albert T.
ASSISTANT EXAMINER: Kilcoyne, John M.

LEGAL REPRESENTATIVE: Behr, Omri M.

NUMEER OF CLAIMS: 10

EXEMPLARY CLAIM: 1

NUMBER OF DERMINGS: 8 Drawing Figure(s): 4 Drawing Page(s)
TAS

LINE COUNT: 786

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method of reducing pathologically excessive levels of activity of collagenolytic enzymes in mammals to substantially normal levels by administering 10-100% of the normal antibiotic therapeutic dose of a tetracycline is disclosed.
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Page 12

=> dup rem 121
PROCESSING COMPLETED FOR L21
L22 19 DUP REM L21 (6 DUPLICATES REMOVED)

=> d ibib ab 1- YOU HAVE REQUESTED DATA FROM 19 ANSWERS - CONTINUE? Y/(N):y

L22 ANSMER 1 OF 19 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 1
ACCESSION NUMBER: 2001:430381 CAPLUS
DOCUMENT NUMBER: 135:164518
Partial characterization of an enzyme fraction with processe activity which converts the spore epptidoglycan hydrolase (SleC) precursor to an active enzyme during germination of Clostridium perfringens \$40 spores and analysis of a gene cluster involved in the activity Shimamoto, Sciko; Moriyama, Ryuichi; Sugimoto, Kazuhiro; Niyata, Shigeru; Makino, Shio
CORPORATE SOURCE: Department of Applied Molecular Biosciences, Graduate School of Bioagricultural Sciences, Nagoya University, Nagoya, 464-8601, Japan Journal of Bacteriology (2001), 183(12), 3742-3751 CODEN: JOBANY: 15SN: 0021-9193 American Society for Microbiology Journal English UAGE: English
A spore cortex-lytic ensyma of Clostridium perfringens
S40 which is encoded by sleC is synthesized at an early stage of
sporulation as a precursor consisting of four domains. After cleavage of
an N-terminal presequence and a C-terminal prosequence during spore
maturation, inactive proenzyme is converted to active enzyme by processing
 of an N-terminal prosequence with germination-specific protease (GSP)
 during germination. The present study was undertaken to characterize In the presence of 3-[(3-cholamidopropyl)dimethyl-ammonio]-1-propanesulfonic acid (CHAPS), a nondenaturing detergent which was needed for the stabilization of GSP, GSP activity was extd. from germinated spores. The enzyme fraction, which was purified to 668-fold by column chromatog., contained three protein components with mol. masses of 60, and 52 kDa. The protease showed optimum activity at pH 5.8 to 8.5 in the presence of 0.18 CHAPS and retained activity after heat treatment at 55.degree. C for 40 min. GSP specifically cleaved the peptide bond

een Val-149 and Val-150 of SleC to generate mature enzyme. Inactivation of GSP by phenylmethylsulfonyl fluoride and HgCl2 indicated that the case is a cysteine-dependent serine protease. Several pieces of evidence demonstrated that three protein components of the enzyme fraction are processed forms of products of cspA, cspB, and cspC, which are positioned in a tandem array just upstream of the 5' end of sleC. The amino acid sequences deduced from the nucleotide sequences of the csp genes showed significant similarity and showed a high degree of homol, with those of the categories the categories considered the categories and the cyanion binding region of subtilisin-like serine proteases. Immunochem, studies suggested that active GSP likely protease

is localized with major cortex-lytic ensymmes on the exterior of the cortex layer in the dormant apore, a location relevant to the pureuit of a cascade of cortex hydrolytic reactions.

REPERENCE COUNT: 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L22 ANSWER 2 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: DOCUMENT NUMBER:

DUS COPYRIGHT 2002 ACS
1999:512145 CAPLUS
131:270291
Human macrophages synthesize type VIII collagen in vitro and in the atherosclerotic plaque
Weitkamp, Benedikt; Cullen, Paul; Plenz, Gabriele;
Robenek, Horst, Rauterberg, Jurgen
Institut fur Arterioskleroseforschung, Munster, AUTHOR (S):

CORPORATE SOURCE: . 48149,

SOURCE:

PUBLISHER:

Germany
FASEB Journal (1999), 13(11), 1445-1457
CODEN: FAJOEC; ISSN: 0892-6638
Federation of American Societies for Experimental
Biology
Journal DOCUMENT TYPE:

MENT TYPE: Journal LAGE: English Type VIII collagen is a short-chain collagen that is present in increased amta. in atherosclerotic lesions. Although the physiol. function of this matrix protein is unclear, recent data suggest an important role in

remodeling. Type VIII collagen in the atherosclerotic lesion is mainly derived from smooth muscle cells. We now show that macrophages in the atherosclerotic vessel wall and monocytes in adjacent mural thrombi also express type VIII collagen. We demonstrated this using a novel combined fluorescence technique that simultaneously stains, within the same tissue section, specific RNAs by in sit thybridization and proteins by indirect immunofluorescence. In culture, human monocyte/macrophages expressed

VIII collagen at all time points from 1 h to 3 wk after isolation. Western blotting and immunopptn. also revealed secretion of type VIII collagen into the medium of 14-day-old macrophages. Because this is first report of secretion of a collagen by macrophage, we tested the effect of lipopolysaccharide (LPS) and interferon .gamma., substances

stimulate macrophages to secrete lytic ensymas, on macrophage expression of type VIII collagen. LPS and interferon .gamma decreased expression of type VIII collagen. By contrast, secretion of matrix metalloproteinase 1 (MMP 1) was increased, indicating a switch

from
a collagen-producing to a degradative phenotype. Double in situ
hybridization studies of expression of type VIII collagen and MMP 1 in
human coronary arteries showed that in regions important for plaque
stability, the ratio of MMP 1 RNA to macrophage type VIII collagen RNA
varies widely, indicating that the transition from one phenotype to the
other that we obsd. in vitro may also occur in vivo.

REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR

FORMAT

RECORD ALL CITATIONS AVAILABLE IN THE RE

L22 ANSWER 1 OF 19 CAPLUS COPYRIGHT 2002 ACS (Continued)

DUPLICATE 1

1

L22 ANSWER 3 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.DUPLICATE

1999:66814 BIOSIS ACCESSION NUMBER: PREV199900066814

DOCUMENT NUMBER: TITLE:

PREVI39900066814
Proteolytic activity of the Gymnorhynchus gigas
plerocercoid: Purification and properties of a collagenase
from the crude extract.
Vazquez-Lopez, C. (1); De Armas-Serra, C.; Gimenez-Pardo,
C.; Rodriguez-Caabeiro, F.
(1) Lab. Parasitol., Fac. Pharm., Univ. Alcala, Alcala de
Henares, Ctra. Madrid-Barcelona Km. 33.600, E-Madrid 28871
Spain AUTHOR(S):

CORPORATE SOURCE:

Henares, Lie., Indexes, Spain
Parasitology Research, (Jan., 1999) Vol. 85, No. 1, pp. 64-70.
ISSN: 0932-0113. SOURCE:

DOCUMENT TYPE: LANGUAGE

MENT TYPE: Article
UNGE: English
The present report demonstrates that the Gymnorhynchus gigas plerocercoid
The present report demonstrates that the Gymnorhynchus gigas plerocercoid
Descesses various types of endo- and exoproteases with activity against
General (azocoil, azocasein, and azoslbumin) and specific substrates
(elastin, keratin, collagen, hemoglobin, fibrinogen, plasma, and
immunoglobulin G). The activity against collagen is principally due to a
24-kDa collagenase with an isoelectric point of 7.5 and without isoforms
or sugar residues. Moreover, its high degree of proteolytic activity
against collagen under conditions similars to those encountered by the
parasite in its hosts (pH and temperature) and its similarity to metalloand cysteine proteases (the principal protease types implicated in
degradation of tissues) suggests the importance of this molecule as a
lytic ensyms principally implicated in penetration
processes across the teleost muscle or/and into the gastrointestinal
system of elasmobranch fishes as well as in molting processes.

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L22 ANSWER 4 OF 19 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1997:316214 CAPLUS
DOCUMENT NUMBER: 127:46817
```

Molecular characterization of a germination-specific muramidase from Clostridium perfringens S40 spores TITLE:

and

AUTHOR (S) :

CORPORATE SOURCE: SOURCE:

PURILI SHER

DOCUMENT TYPE: LANGUAGE:

Molecular characterization of a germination-specific muramidase from Clostridium perfringens \$40 spores

nucleotide sequence of the corresponding gene Chen. Yinghus: Miyata, Shigeru; Makino, Shio; Moriyama, Ryuichi
Dep. Applied Biol. Sci., Sch. Agric. Sci., Nagoya Univ., Nagoya-hchi, 464-01. Japan
Journal of Bacteriology (1997), 179(10), 3181-3187 CODEN: JOBANY; ISSN: 0021-9183
American Society for Microbiology
MENT TYPE: Journal
SUAGE: Host Sci., Sch. Agric. Sci., Nagoya Univ., Nagoya-hchi, 464-01. Japan
JOURNAL TYPE: Journal
SUAGE: Host Microbiology
MENT TYPE: Journal
SUAGE: Host Microbiology
MENT TYPE: Journal
SUAGE: Host Microbiology
MINT TYPE: Journal
SUAGE: Host Microbiology
MINT TYPE: Journal
SUAGE: Host Microbiology
MINT TYPE: Journal
SUAGE: Host Miryahara, and S. Makino, Microbiol. 141:2643-2650, 1995). The lytic ensyma was purified to homogeneity by
anion-exchange chromatog. and shown to be a muramidase which requires
divalent cations (Ca2+, Mg2+, or Nn2+) for its activity. The enzyme was
inactivated by sulfhydryl reagenta, and sodium thioglycolate reversed the
inactivation by Hg2+. The muramidase hydrolyzed isolated appore cortical
fragments from a variety of wild-type organisms but had minual activity
on decoated appores and isolated cell walls. However, the enzyme was not
capable of digesting isolated cortical fragments from spores of Bacillus
subtilis ADDI, which lacks muramic acid delta-lactam in its cortical
preptidoglycan. This indicates that the enzyme recognizes the
delta-lactam residue peculiar to spore peptidoglycan, suggesting an
involvement of the enzyme in spore germination. Immunochem. studies
indicated that the muramidase in its mature form is localized on the
muramidase, aleM, was cloned into Escherichia coli, and the nucleotide
sequence was detd. The gene encoded a protein of 221 amino acids with a
deduced mol. wt. of 36,358. The deduced amino acid sequence of the sleM
gene indicated that the muramidase belongs to a sep. group within the lysozyme
family typified by the fungus Chalaro

L22 ANSWER 6 OF 19 USPATFULL SCREENIN NUMBER: 96:388

PATFULL
96:38811 USPATFULL
DNA fragment encoding a hydrogen peroxide-generating NADH oxidase

INVENTOR (S): Higuchi, Masako, Neyagawa, Japan Higueni, Mesako, Neyagawa, Japan Mataumoto, Junichi, Moriguchi, Japan Yamamoto, Yoshikazu, Neyagawa, Japan Kamio, Yoshiyuki, Sendai, Japan Izaki, Kazuo, Miyagi, Japan Nippon Paint Co., Ltd., Osaka, Japan (non-U.S.

PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE PATENT INFORMATION: APPLICATION INFO.: US 5514587 US 1994-220677 19960507 19940331 (8)

NUMBER DATE PRICRITY INFORMATION: 19930331

JP 1993-73989 JP 1993-254459 Utility Granted DOCUMENT TYPE: FILE SEGMENT: PRIMARY EXAMINER: ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: NUMBER OF CLAIMS: EXEMPLARY CLAIM: Wax, Robert A. Hobba, Lisa J. Townsend & Banta

LINE COURT: 985
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB A gene encoding a hydrogen peroxide-generating NADH oxidase and a method

for preparing a large amount of the NADH oxidaae with the use of the gene and gene recombinant techniques are diaclosed.

19931012

L22 ANSWER 5 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: DOCUMENT NUMBER: 1997:358545 CAPLUS

126:329591

Production and wound-healing applications of enzyme preparations obtained from lysis of Streptomyces TITLE: flavus 197

INVENTOR(S):

flavus 197 Grigiskis, Saulius; Spokiene, Aldona-Ona; Baskys, Egidijus-Vladas; Vilutis, Kestutis Grigiskis, Saulius, Lithuanis; Spokiene, Aldona-Ona; Baskys, Egidijus-Vladas; Vilutis, Kestutis Lith., 7 pp. CODEN: LIXXFS PATENT ASSIGNEE(S):

SOURCE:

DOCUMENT TYPE: Patent Lithuanian ANGUAGE

PAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

LT 1878 B 19960425 LT 1994-1868 19940202
Enzymes can be obtained by lysis of cells of Streptomyces flavus 197
(deposited at the UAB Biocenter microorganism collection under the accession no. K-91) grown in liq. culture by filtration, concn., and

extn.

of the dry enzyme complex with org. solventa. The enzyme prepn. obtained is a complex of lytic enzymes, in particular proteolytic peptidases, collagenases, and elastases which can be effective for lymis of some gram-pos. and gram-neg. microorganisms, including pathogens. The enzyme prepns. have medical and veterinary applications in

treatment of purulent and necrotic wounds, bed sores, burns, and bullet wounds.

L22 ANSWER 7 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. DUPLICATE

ACCESSION NUMBER: 1996:505492 BIOSIS PREV199699227848

DOCUMENT NUMBER: TITLE:

Fractions from commercial collagenase preparations: Use in enzymic isolation of the islets of Langerhans from porcine

pancreas. Kloeck, Gerd; Kowalski, Matthias B.; Herring, Bernhard J.; Eiden, Martin E.; Weidemann, Astrid; Langer, Stefan; Zimmermann, Ulrich (1); Pederlin, Konrad; Bretzel, AUTHOR (S):

Reinhard

CORPORATE SOURCE:

SOURCE -

DOCUMENT TYPE:

G.

ORATE SOURCE: (1) Lehratuhl Biotechnologie, Univ. Wuerzburg, Biozentrum,
Am Hubland, D-97074 Wuerzburg Germany
CE: Cell Transplantation, (1996) Vol. 5, No. 5, pp. 543-551.
ISSN: 0983-6897.
MENT TYPE: Article
UMGE: English
Transplantation of isolated islets of Langerhans is an intriguing
possibility for the treatment of diabetes mellitus. The isolation of
islets from pancreata requires the apecific dissociation of the tissue.
Commercial collagenases from Clostridium histolyticum are widely used for
this purpose. Unfortunately, the effectiveness of these commercial
mes

this purpose. Unfortunately, the effectiveness of these commercial mes is not predictable and differs considerably between suppliers and even from lot to lot. This is due mainly to differences in their specific collagenase activity and to the presence of other lytic marymas, as well as to other contaminants. Pree flow zone electrophoresis (PPZE) was used to separate the effective protein components from undesired compounds and to prepare a digestive enzyme mixture with controlled composition of lytic activities. Practionation of crude collagenases by PPZE resulted in partially purified protein fractions that were enriched for collagenase and tryptic activities, and contained only trace amounts of neutral protease. These preparations proved to be highly effective in an in vitro assay for the liberation of viable islets from porcine pancreas. To scale up the production of these collagenases with defined enzyme composition, we fractionated two different lots of a commercial collagenase from C. histolyticum (one lot effective in islet isolation, the other not) by using fast protein liquid chromatography (FPLC) on hydroxyapatite. Again, high efficacy of islet release from pancreast cissue was correlated to high specific tryptic

collagenase activities and low levels of neutral protesse. The chromatographic protocol developed in this study converted a effective collagenase lot into a preparation that allowed successful islet isolation.

L22 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1995:900763 CAPLUS DOCUMENT NUMBER: 124:47021

124:47021 A gene (sleC) encoding a spore-cortex-lytic ensymm from Clostridium perfringes S40 spores; cloning, sequence analysis and molecular characterization

AUTHOR (S):

Miyata, Shigeru; Moriyama, Ryuichi; Miyhara, Nobuko; Makino, Shio CORPORATE SOURCE:

SOURCE:

PUBLISHER: CUMENT TYPE:

LANGUAGE

ON(5): Mayara, Shigeru; Moriyama, Kyulchi; Miyhara, Nobuko;
Makino, Shigicultural Sciences, Nagoya University,
Alchi, 46-01, Japan
CE: Microbiology (Reading, United Kingdom) (1995),
141(10), 2643-50
CODEN: MROBED; ISSN: 1350-0872
JOHER: Society for General Microbiology
MENT TYPE: Journal
LUNGE: English
Antiserum was raised against a 31 kDa spore-cortex-lytic
ensyma, which is released during germination of Clostridium
perfringens S40 spores. Western blotting of dormant spore and vegetative
cell fractions sept by SDS-PAGE indicated that the 31 kDa enzyme is
spore-specific and that the enzyme in the dormant spore exists as a 36

protein which has no cortex-lytic activity. A gene encoding the 31 kDa enzyme, slec, was cloned into Escherichia coli using a synthetic oligonucleotide as a hybridization probe and the nucleotide sequence of the entire gene was detd. The N-terminal amino acid sequence of the 36 kDa protein was found in this reading frame, confirming that the 36 kDa protein is a pro-form of the 31 kDa enzyme. The deduced amino acid sequence indicated that the 31 kDa enzyme is produced as a precursor, comprising three portions; an N-terminal prepro-sequence (114 amino acid residues), a pro-sequence (35 amino acid residues) and a mature enzyme (289 amino acid residues). It is suggested that the 36 kDa pro-enzyme is non-covalently attached to the exterior of the cortex layer, and that the pro-form is processed to release the active enzyme during germination.

L22 ANSWER 10 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1994:652315 CAPLUS

1994:652315 CAPLUS 121:252315 DOCUMENT NUMBER:

TITLE:

Correlation of proteolytic activities of organ cultured intact mouse skin with defined hair cycle

stages AUTHOR (S):

stages
Paus, Ralf; Krejci-Papa, Niels; Li, Lingna;
Czarnetzki, Beate M.; Hoffman, Robert M.
Dept. Dermatology, University Hospital R. Virchow,
Berlin, D-13344, Germany
J. Dermatol. Sci. (1994), 7(3), 202-9
CODEN: JDSCEI; ISSN: 0923-1811 CORPORATE SOURCE:

SOURCE:

DOCUMENT TYPE: Journal English

The cyclic growth activity of the hair follicle is characterized by substantial remodelling of the extracellular matrix, yet, little is known about the proteolytic activities regulating this process. In murine

hair cycling is highly synchronized and is assocd. With dramatic remodelling of all skin compartments. The authors therefore assessed, in this pilot study, proteolytic activities of murine skin from various stages of the depilation-induced hair cycle. The defined proteolytic activities displayed by organ cultured intact mouse skin differed between hair cycle stages. Skin with all follicles in telogen or mid anagen displayed only minimal lysis of collagen type I gels, while early anagen skin had significant collagenase activity. Skin cultured on gelatin gels at the air-liq. interphase (histoculture) completely lysed the gel within 5 days when all follicles were in early anagen, whereas this was not obsd. With mid and very late anagen skin. Zymog, of conditioned medium from these cultures revealed the secretion of activated interstitial collagenase and of gelatinases of 72 and 92 kDa, with the max. of interstitial collagenase activity secreted by anagen IV skin. Addn. of TPA or TNP-.elpha. to the culture medium stimulated secreted collagenase type I activity. The C 57 BL-6 mouse offers an attractive model for dissecting and manipulating hair cycle-assocd. proteolysis in a physiol. relevant system.

L22 ANSWER 9 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1995:366297 CAPLUS

DOCUMENT NUMBER: 122:180018

Tacking the evolution of the bacterial choline-binding domain: molecular characterization of the Clostridium acetobutylicum NCIB 8052 cspA gene Sanchez-Beato, Ana R.; Ronda, Concepcion; Garcia, AUTHOR (S):

CORPORATE SOURCE: Consejo Superior Investigaciones Cientificas, Madrid,

Spain Journal of Bacteriology (1995), 177(4), 1098-103 CODEN: JOBAAY; ISSN: 0021-9193 American Society for Microbiology SOURCE:

PUBLISHER:

DOCUMENT TYPE: LANGUAGE:

AGE: English
The major secreted protein of Clostridium acetobutylicum NCIB 8052, a choline-contg. strain, is CspA (clostridial secreted protein). It

ars
to be a 115,000-Mr glycoprotein that specifically recognizes the choline
residues of the cell wall. Polyclonal antibodies raised against CspA
detected the presence of the protein in the cell envelope and in the
culture medium. The sol. CspA protein has been purified, and an
oligonucleotide probe, prepd. from the detd. N-terminal sequence, has

used to clone the cspA gene which encodes a protein with 590 amino acids and an Mr of 63,740. According to the predicted amino acid sequence,

is synthesized with an N-terminal segment of 26 amino acids acteristic of prokaryotic signal peptides. Expression of the cspA gene in Escherichia coli led to the prodn. of a major anti-CspA-labeled protein

80,000 Da which was purified by affinity chromatog. on DEAE-cellulose. A comparison of CspA with other proteins in the EMBL database revealed that the C-terminal half of CspA is homologous to the choline-binding domains of the major pneumococcal autolysin (LytA amidsse), the pneumococcal antigen PspA, and other cell wall-lytic enzymas of pneumococcal phages. This region, which is constructed of four repeating motifs, also displays a high similarity with the glucan-binding domains

several streptococcal glycosyltransferases and the toxins of Clostridium difficile.

L22 ANSWER 11 OF 19 USPATFULL

CCESSION NUMBER

SPATFULL

93.46351 USPATFULL

Device and method for the rapid qualitative and quantitative determination of the presence of a reactive ligand in a fluid

Marchand, Joseph, Verrieres Le Buisson, France

Toledano, Jacques, Paris, France

Compagnie Oris Industrie S.A., Gif-Sur-Yvette, France (non-U.S. corporation)

Cistest, Paris, France (non-U.S. corporation) TITLE:

INVENTOR(S):

PATENT ASSIGNEE(S):

MBER KIND NUMBER DATE PATENT INFORMATION: US 5217905 US 1988-220895 19930608 APPLICATION INFO. : 19880718 (7)

NUMBER DATE

PRIORITY INFORMATION: DOCUMENT TYPE: PILE SEGMENT: 19880428

FR 1988-5668 1 Utility Granted Saunders, David Chin, Christopher L. Browdy and Neimark PRIMARY EXAMINER ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: NUMBER OF CLAIMS: EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

4 Drawing Figure(s); 3 Drawing Page(s) LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A device for the rapid qualitative and quantitative determination of

presence of a reactive ligand in a fluid.

This device comprises a first reaction zone in which there is an at least temporarily impermeable membrane designed to receive a sample of test fluid and to be associated with at least one labeled reagent; a second reaction zone which is bounded on the one hand by the said membrane and on the other by a second at least temporarily impermeable membrane comprising a solid phase containing a reference reagent; and a third reaction zone which contains means for developing the reaction.

A method for the rapid qualitative and quantitative determination of the

presence of a reactive ligand in a fluid.

Applications to the detection of the presence, in a biological fluid,

antibodies or antigens in particular.

L22 ANSWER 12 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1993:487507 CAPLUS DOCUMENT NUMBER: 119:87507 DUPLICATE 4

119:87507

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AUTHOR (5):

CORPORATE SOURCE:

SOURCE:

DOCUMENT TYPE: LANGUAGE:

CODEN: GENEDS; ISSN: 0378-1119

JOURNAL
JOURNAL
JOURNAL
JOURNAL
The lyc gene, encoding an autolytic lysozyme from C. acetobutylicum
ATCC824, has been cloned. The nucleotide sequence of the lyc gene has
been detd. and found to encode a protein of 324 amino acids (aa) with a
deduced Mr of 34,939. The lyc gene is preceded by 2 open reading frames
with unknown functions, suggesting that this gene is part of an operon.
Comparison between the deduced as sequence of the lyc gene and the
directly detd. N-terminal sequence of the extracellular clostridial
lysozyme suggests that the enzyme is synthesized without a cleavable
signal peptide. Moreover, the comparative analyses between the
clostridial lysozyme and other known cell-wall lytic
susymas revealed a significant similarity with the N-terminal
portion of the lysozymes of Streptomyces globisporus, the fungus
Chalaropsis, the Lactobacillus bulgaricus bacteriophage mvi, and the
Streptomyces pneumoniae bacteriophages of the Cp family (CPL lysozymes).
In addn., the analyses showed that the C-terminal half of the clostridial
lysozyme was homologous to the N-terminal domain of the
muramoyl-pentapeptide-carboxypeptidase of Streptomyces albus, suggesting

role in substrate binding. The existence of 5 putative repeated motifs

the C-terminal region of the autolytic lysozyme suggests that this region could play a role in the recognition of the polymeric substrate.

L22 ANSWER 13 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1986:49379 CAPLUS DOCUMENT NUMBER: 104:49379

Invasion potential of human choriocarcinoma cell TITLE:

lines

and the role of lytic ensymmes
Sekiya, S.; Ocsaki, T.; Suzuki, N.; Takamizawa, H.
Sch. Med., Chiba Univ., Chiba, 280, Japan
Gynecol. Oncol. (1985), 22(3), 324-33
CODEN: GYNQA3; ISSN: 0090-8258 AUTHOR (5) : CORPORATE SOURCE:

DOCUMENT TYPE: LANGUAGE:

CODEN: GYNOAJ; ISSN: 0090-8258

MENT TYPE: Journal

UAGE: English

The chick choricallentoic membrane (CAM) was used as an assay system to examine the invasive potential of human choriccarcinoma cell lines. When 5 times. 106 cells were inoculated into the CAM at the 10th day of postfertilization. 3 of 8 cell lines formed extensively invasive cumors within the CAM. The tumorigenic potential of cell lines in hammater chek pouches was correlated with their invasive potential in the CAM. The invasive capacity of cell lines correlated well with the amt. of collagenase but did not correlate with the amt. of plasminogen activator or cathepsin B secreted by them.

L22 ANSWER 14 OP 19 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1983:486054 CAPLUS
ODCUMENT NUMBER: 99:86054
CTITLE: Changes of the activity of lytic
emsymmes in alveolar bone incident to
orthodontic tooth movement
Kamei, Teruaki
CORPORATE SOURCE: Dep. Orthod, Kanagawa Dent. Coll., Yokosuka, Japan
SOURCE: CODEN: KSHGDM
DOCUMENT TYPE:

CODEN: KSHGDM

DOCUMENT TYPE: Journal
LANGUAGE: Japanese
AB Helical torsion spring which exerted about 50g expanding force was
attached to the maxillary incisors of rabbita. After 1, 2, 4, 7, 14, or
21 days of tooth movement, the animals were sacrificed and the alveolar
bones assayed for 6 enzymes. Owing to the helical torsion spring, the
right and left upper incisors sepd. rapidly, and tooth movement
continued.
The orthodontic tooth movement differed according to 3 stages. Following

The orthodontic tooth movement differed according to 3 stages. Follothe initial tooth displacement, many enzyme activities were elevated

the control level, which suggested that metabolic changes in the alveolar bone cells were induced by orthodontic stimulation (first stage). Cathepsin D and collagenase activities increased after tooth movement and reached their peak levels at days 2-4 (second stage), but afterwards they gradually increased following orthodontic tooth movement until days 7-14 (second stage). Alk. phosphatase and cathepsin B activities gradually increased following orthodontic tooth movement until days 7-14 (second stage). Alk. phosphatase and hyaluconidase activities rapidly increased on the 2nd day after tooth movement, and continued to increase to day 21 (third stage). Apparently, resorption of alveolar bone shifted to bone formation at day 7 of orthodontic tooth movement, and after that time an active bone remodeling took place.

L22 ANSWER 15 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1979:52082 CAPLUS

90:52082 A histochemical study about the influence of DOCUMENT NUMBER: TITLE:

lytic ensymmes on plasma membrane enzyme activities in rat liver and kidney Hardonk, M. J.; Meskendorp-Haarsma, T. J.; Koudstaal,

AUTHOR (S):

J.
Dep. Pathol., Univ. Groningen, Groningen, Neth.
Histochemistry (1978), 58(3), 177-81
CODEN: HCMYAL; ISSN: 0301-5564 CORPORATE SOURCE: SOURCE:

DOCUMENT TYPE: Journal English

LINGE.

English

The effect of lipolytic, glycolytic, and proteolytic enzymes on the activities of plasma membrane enzyme activities in rat liver and kidney was investigated by a pretreatment of tissue sections with the lytic ensymas. The action of proteolytic enzymes causes a very strong decrease of leucyl-.beta.-naphthylamidase activity, whereas the activities of ATPase, S'-nucleotidase, and alk, phosphatase show a lesser degree. This indicates a different membrane anchorage of leucyl-.beta.-naphthylamidase as compared to that of the phosphateases. Treatment with glycolytic enzymes results in a decrease of uncleotidase.

Treatment with glycolytic enzymes results in a decrease of 5-nucleotidase and ATPase activity, whereas liver alk, phosphatase and leucyl-beta-naphthylamidase show an increase in activity. Treatment with phospholipase C gives about the same results. The very strong decrease of 5-nucleotidase activity indicates a great dependence on

L22 ANSWER 16 OF 19 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER:
DOCUMENT NUMBER:
1978:102967 CAPLUS
88:102967
Proliferative synovitis in hemophilis. Biochemical and morphologic observations
AUTHOR(S):
Mainardi, Carlo L.; Levine, Peter H.; Werb, Zena; Harris, Edward D., Jr.
CORPORATE SOURCE:
Dep. Med., Dartmouth-Hitchcock Med. Cent., Hanover, N.

CORPORATE SOURCE:

SOURCE:

DOCUMENT TYPE:

H., USA

CE: Arthritis Rheum. (1978), 21(1), 137-44

CODEN: ARHEAW; ISSN: 0004-3591

MENT TYPE: Journal

UNGS: English

The synovium removed from the knee of a 10-yr-old with hemophiia A was characterized morphol. and biochem. The specimen showed villous hypertrophy with hyperplasia of synovial lining cells which contained abundant intracytoplasmic granules of hemosiderin. Monolayer cultures prepd. from enzymically dispersed tissue were characterized by pigment-laden fibroblast-like cells and round cells. Both explants of synovium and adherent cells secreted a large amt. of latent collagenase and neutral proteinase into the culture medium. The secretion of these enzymes dropped sharply and intracellular pigment decreased with passage of these cultures. Lysozyme was secreted by the explants but was not detected in the monolayer culture medium. These data establish the degradative potential of the synovitis found in hemophilia and support

concept that recurrent hemoarthrosis without inflammation is sufficient

and of itself to produce synovitis.

L22 ANSWER 18 OF 19 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1976:144611 CAPLUS

DOCUMENT NUMBER : 84:144611 TITLE:

vitamin

Inhibition of plasmin-mediated fibrinolysis by

AUTHOR (S)

Moroz, L. A.; Gilmore, N. J. McGill Univ. Clin., R. Victoria Hosp., Montreal, CORPORATE SOURCE:

SOURCE:

Can.
Nature (London) (1976), 259(5540), 235-7
CODEN: NATUAS
Journal

DOCUMENT TYPE: LANGUAGE:

MENT TYPE: Journal MAGE: English English In vitro fibrinolysis mediated by plasmin [9001-90-5], but not by other lytic ensymmas, e.g. collagenase [9001-12-1], was inhibited by vitamin E [1405-18-4] or D. alpha.-tocopheryl succinate [4345-03-3], 50% inhibition occurring at an inhibitor:enzyme molar ratio of .apprx.] 50% inheritor occurring at an inhibitor:enzyme molar ratio of .apprx.] 100:1. The pertinence of these data to physiol. and pathol. phenomena, and to vitamin E therapy is discussed.

L22 ANSMER 17 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. ACCESSION NUMBER: 1976:179611 BIOSIS DOCUMENT NUMBER: BA62:9611

MOLECULAR FORMS OF ACETYL CHOLIN ESTERASE FROM TORPEDO-CALIFORNICA THEIR RELATIONSHIP TO SYNAPTIC TITLE:

MEMBRANES.

LWEBUGA-MUKASA J S: LAPPI S: TAYLOR AUTHOR(S): AUTHORIS: DEBUGN-RUKASA J S; LAPPI S; IATLOR F S SCURCE: BICHAMN. 15SN: 0006-2960.

FILE SEGMENT: BA; OLD
LANGUAGE: Unavailable
AB The 16S and 85 forms of acetylcholinesterase (AchE), which are composed

an elongated tail structure in addition to the more globular catalytic subunits, were extracted and purified from membranes from T. californica electric organs. Their subunit compositions and quaternary structures

compared with 11S lytic ensyme which is derived from collagenase or trypsin treatment of the membranes and devoid of the tail unit. Upon sodium dodecyl sulfate-polyacrylamide gel electrophoresis in the absence of reducing agent, appreciable populations of monomeric through tetrameric species were observed for the 11S form. Under the same conditions the 18S form yielded only monomer and dimer in addition to a higher molecular weight species. If complete reduction was effected, only the 80,000 molecular weight monomer was dominant for both the 11S and 16S forms. Cross-linking of the 11S form by dimethyl suberimidate followed by reduction yielded monomer through tetramer in descending frequency, while the 16S form again showed a high molecular weight species. A comparison

the composition of the 11S and 16S forms revealed that the latter had an increased glycine content, and 1.1 and 0.3 mol % hydroxyproline and hydroxylysine, respectively. Collagenases that were purified to homogeneity and were devoid of amidase and caseinolytic activity, but active against native collagen, converted 16S acetylcholinesterase to the 11S form. Composition and substrate behavior of the 16S enzyme were indicative of the tail unit containing a collagen-like sequence. A membrane fraction enriched in AchE and components of basement membrane were separated from the major portion of the membrane protein. The 16S

not the 11S form reassociated selectively with this membrane fraction. There are distinct similarities between the tail unit of ${\sf AchE}$ and

basement membrane components, and there may be a primary association of AchE with the basement membrane.

L22 ANSWER 19 OF 19 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. ACCESSION NUMBER: 1976:34318 BIOSIS

1976:34318 BIOSIS BR12:34318

DOCUMENT NUMBER: TITLE:

BRI2:34318
TRYPAN BLUE INHIBITION OF LYTIC EMETHE
ACTIVITY DURING NEWT LIMB REGENERATION.
DEARLOWS G E; DRESDEN N H
J. Cell Biol., (1975) 67 (2 PART 2), 87A.
CODDEN JCLEBA3. ISSN: 0021-9525.
CONFERENCE
BR; OLD
Unavailable AUTHOR(S):

DOCUMENT TYPE: FILE SEGMENT: LANGUAGE: